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Sheet 1 of 3. Form PTO-1449 U.S. Department of Commerce ATTY. DOCKET NO. INTERNATIONAL APPLICATION NO. Patent and Trademark Office PB60213 PCT/EP04/04001 <u> 10/552571</u> INFORMATION DISCLOSURE STATEMENT APPLICANT Barry BARTON, Susan JENSEN, Alison Michelle BY APPLICANT GRIFFIN and Annie WONG FILING DATE **GROUP** (Use several sheets if necessary) Herewith **U.S. PATENT DOCUMENTS** Examiner Document Date Name Class Subclass Filing Date Initial Number If Appropriate AA4,202,819 5/13/80 Kellett et al. 260 245.3 FOREIGN PATENT DOCUMENTS Document Date Country Class <u>Translation</u> Subclass Number Yes No BAWO98/33896 06 Aug 1998 **PCT** BBWO01/30977 03 May 2001 **PCT** BCWO03/040372 15 May 2003 **PCT** BDWO00/03581 27 Jan. 2000 **PCT** BE JP 53-104796 12 Sept 1978 Japan BFES 550549 1987 Spain BGEP 0 550549 20 Sept 1990 EP BH 2,108,113 10 Apr 1995 Canada OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.) Liras P., et al., "Clavulanic acid, a ß-lactamase inhibitor: biosynthesis and molecular CA genetics," Applied Microbiology and Biotechnology, Springer Verlag Berlin, Germany (2000) 54(4):467-475. Mosher R., et al., "Genes Specific for the Biosynthesis of Clavam Metabolites Antipodal to CB Clavulanic Acid are Clustered with the Gene for Clavaminate Synthase 1 in Streptomyces Clavuligerus," Antimicrobial Agents and Chemotherapy, American Society for Microbiology, Washington, DC (May 1999) 43(5):1215-1224. A.L. Demain, "Biosynthesis and Regulation of Beta-Lactam Antibiotics", 50 Years of CC Penicillin Applications, History & Trends (1990). Townsend, et al., "Biosynthesis of clavulanic Acid: Origin of the C₃ Unit," J. Am. Chem. Soc., CD 107(4):1066-1068 (1985). Valentine, et al., "Evidence that Arginine is a Later Metabolic Intermediate than Ornithine in CE the Biosyntheis of Clavulanic Acid by Streptomyces Clavuligerus," J. Chem. Soc. Chem. Comm., 15:1210-1211 (1993). **EXAMINER** DATE CONSIDERED /Scott Long/ 04/11/2008

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		GRIFFIN and Annie WONG		
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OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

CF	Janc, et al., "Emerging Evidence for a Shared Biosynthetic Pathway Among Clavulanic Acid		
	and the Structurally Diverse Clavam Metabolites," <i>Bioorg. Med. Chem.</i> Lett., 3:2313-2316 (1993).		
CG	Rohl, et al., "Biological properties and mode of action of clavams," Arch. Microbiol., 147:315-320 (1987).		
СН	Aidoo, et al., "Cloning, sequencing and disruption of a gene from Streptomyces clavuligerus involved in clavulanic acid biosynthesis," Gene, 147:41-46 (1994).		
CI	Sambrook, et al., <i>Molecular Cloning: A Laboratory Manual</i> , 2nd Edition. Cold Spring Harbor Laboratory, Cold Spring Harbor, NY (1989).		
CJ	"Postassium Clavulanate," British Pharmacopoeia (1993); Addendum (1994) pp. 1362-1363.		
CK	"Clavulanate Potassium," British Pharmacopoeia Official Monographs, USP 23 NF18 pp: 384-385 (1985).		
CL	Hopwood, et al., Genetic Manipulation of Streptomyces. A Cloning Manual (1985).		
CM	Stahl, et al., "Development and Application of Nucleic Acid Probes," <i>Nucleic Acid Techniques in Bacterial Systematics</i> . Ed. E. Stackebrandt & M. Goodfellow. Toronto: John Wiley & Sons, pp. 204-248 (1991).		
CN	Doran, et al., "Isolation and Characterization of a \(\beta\)-Lactamase-Inhibitory Protein from Streptomyces Clavuligerus and Cloning and Analysis of the Corresponding Gene," J. Bacteriol. 172(9):4909-4918 (1990).		
СО	Vieira, et al., "Production of Single-Stranded Plasmid DNA," Methods Enzymol., 153: 3-11 (1987).		
СР	Sanger, et al., "DNA Sequencing with Chain-terminating Inhibitors," <i>Proc. Natl. Acad. Sci. USA</i> , 74: 5463-5467 (1977).		
CQ	Ward, et al., "Construction and Characterisation of a Series of Multi-Copy Promoter-Probe Plasmid Vectors for Streptomyces Using the Aminoglycoside Phosphotransferase Gene from Tn5 as Indicator," Mol. Gen. Genet. 203: 468-478 (1986).		
CR	Pruess, et al.,"A New Clavam Antibiotic from Streptomyces Clavuligerus," <i>Journal of Antibiotics</i> , XXXVI(3): 208-212 (Mar. 1983).		
CS	Paradkar, et al., "Functional analysis of the Gene Encoding the Clavaminate Synthase 2 Isoenzyme Involved in Clavulanic Acid Biosynthesis in Streptomyces Clavuligerus," <i>Journal of Bacteriology</i> , 177: 1307-1314 (1995).		
СТ	Hodgson, J.E. et al., "Clavulanic Acid Biosynthesis in Streptomyces Clavuligerus: Gene Cloning and Characterization," <i>Gene</i> , (1995) Vol. 166, pp. 49-55		
EXAMINER	/Scott Long/ DATE CONSIDERED 04/11/2008		

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Barry BARTON, Susan JENSEN, Alison Michelle INFORMATION DISCLOSURE STATEMENT BY APPLICANT GRIFFIN and Annie WONG GROUP FILING DATE Herewith (Use several sheets if necessary)

CU
Busby, R.W., "Expression and Purification of Two Isozymes of Clavaminate Synthase and Initial Characterization of the Iron Binding Site," J. Biological Chem., (1995), Vol. 270(9), pp. 4262-69

CV
Brown, D. et al., "Structures of Three Novel \$\beta\$-lactams Isolated from Streptomyces Clavuligerus," J.C.S. Chem. Comm., (1979), pp. 282-3

CW

Formation: Cloning and Sequencing of Both Genes from Streptomycs Clavuligerus," Biochemistry, (1992), Vol. 31, pp. 12648-57 Elson, et al., "Studies on the Biosynthesis of Clavulanic Acid." J. Antibiotics, XXXI(6): 568 CXEvans, R. H. et al., "Ro 22-5417, A New Clavam Antibiotic from Streptomyces Clavuligerus CY II. Fermentation, Isolation and Structure" J. Antiobiotics, (1983), Vol. 36(3), pp. 213-216 Muller, J-C. et al, "Ro 22-5417, A New Clavam Antibiotic from Streptomyces Clavuligerus CZIII Absolute Stereochemistry," J. Antiobiotics, (1983), Vol. 36(3), pp. 216-224 King, H.D. et al., "Clavamycins, New Clavam Antibiotics from Two Variants of CAA Streptomyces Hygroscopicus I. Taxonomy of the Producing Organisms, Germentation, and Biological Activities," J. Antiobiotics, (1986), Vol. 39(4), pp. 510-15 Janc, J.W. et al., "Purification and Characterizatgion of Clavaminate Synthase from CBB Streptomyces Antibioticus," J. Biological Chem., (1995), Vol. 270(10), pp. 5399-5404 Iwata-Reuly, D. and C. A. Townsend, "Common Origin of Clavulanic Acid and Other

Marsh, E.N. et al., "Two Isozymes of Clavaminate Synthase Central to Clavulanic Acid

Streptomyces Antibioticus," J. Biological Chem., (1995), Vol. 270(10), pp. 5399-5404

CCC Iwata-Reuly, D. and C. A. Townsend, "Common Origin of Clavulanic Acid and Other Clavam Metabolites in Streptomyces," J. Am. Chem. Soc., (1992), Vol. 114, pp. 2762-3

CDD Paradkar, A. S. and S. E. Jensen, "Functional Analysis of the Gene Encoding the Clavaminate Synthase 2 Isoenzyme Involved in Clavulanic Acid Biosynthesis in Streptomyces Clavuligerus," J. Bacteriology, (1995), Vol 177(5), pp. 1307-14

CEE Baldwin, J. E. et al., "Enzymes of Valcalavam Biosynthesis," Tetrahedron Letters, (1994), Vol. 35(17), pp. 2783-6

CFF Egan, L.A. et al., "Probable Role of Clavaminic Acid as the Terminal Intermediate in the Common Pathway to Clavulanic Acid and the Antipodal Clavam Metabolites," J. Am. Chem. Soc., (1997), Vol. 119, pp. 2348-55

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